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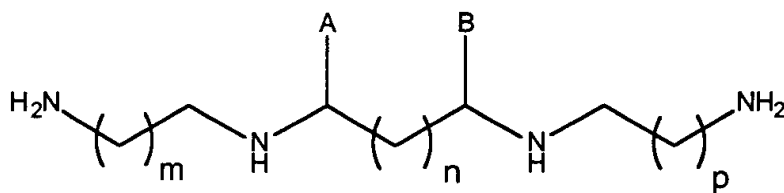
## Claims

We claim;

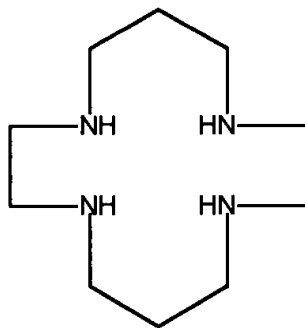
- 1 1 A method of treating degenerative diseases due to  
2 acquired mitochondrial DNA damage  
3 redox damage to mitochondrial macromolecules  
4 and inherited mitochondrial genetic defects  
5 said method comprising the steps of: selecting a composition from a group consisting of open  
6 ring polyamines, macrocyclic polyamines, branched linear polyamines and substituted  
7 polyamines;  
8 synthesizing said composition; and  
9 administering an effective dose of said composition to a mammal.

process of use  
by process  
of making

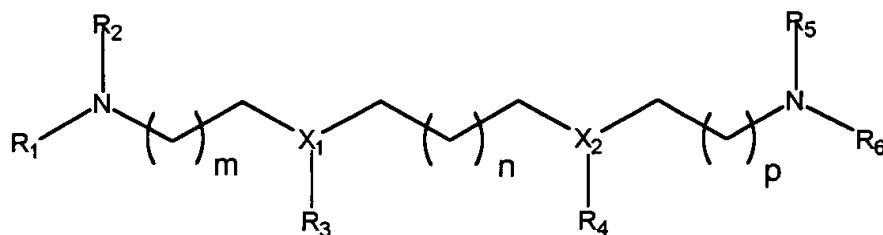
- 1 2 The method of Claim 1 wherein said step of synthesizing comprises converting by treatment  
2 with an alkyl halide a compound taken from a group consisting of those compounds having the  
3 formula



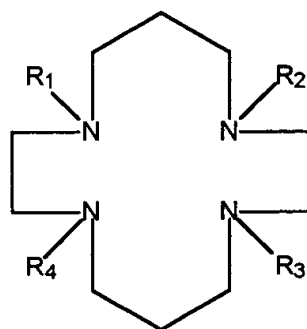
7  
8 wherein A and B are hydrogen or alkyl, and m, n, and p are the same or different, and those  
9 compounds having the formula



3 The method of claim 2 wherein said composition is taken from a group consisting of those compositions having the formulae:



and



wherein:

18 R<sub>1</sub> and R<sub>2</sub> are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid,  
19 glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol,  
20 vitamin E, hydroxytoluene, carvidilol, α-lipoic acid, α-tocopherol, ubiquinone,  
21 phylloquinone, β-carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme  
22 Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene, –  
23 (CH<sub>2</sub>)<sub>n</sub>[XCH<sub>2</sub>]<sub>n</sub>NH<sub>2</sub> - wherein n = 3-6 and R<sub>1</sub> and R<sub>2</sub> taken together are –(CH<sub>2</sub>XCH<sub>2</sub>)<sub>n</sub>–  
24 wherein n = 3-6,

25 R<sub>3</sub> and R<sub>4</sub> are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid,  
26 glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol,  
27 vitamin E, hydroxytoluene, carvidilol, α-lipoic acid, α-tocopherol, ubiquinone,  
28 phylloquinone, β-carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme  
29 Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene or  
30 heterocycle and R<sub>3</sub> and R<sub>4</sub> taken together are –(CH<sub>2</sub>XCH<sub>2</sub>)<sub>n</sub>– wherein n = 3-6,

31 R<sub>5</sub> and R<sub>6</sub> are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid,  
32 glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol,  
33 vitamin E, hydroxytoluene, carvidilol, α-lipoic acid, α-tocopherol, ubiquinone,  
34 phylloquinone, β-carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme  
35 Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene –  
36 (CH<sub>2</sub>)<sub>n</sub>[XCH<sub>2</sub>]<sub>n</sub>NH<sub>2</sub> - wherein n = 3-6, and R<sub>5</sub> and R<sub>6</sub> taken together are –(CH<sub>2</sub>XCH<sub>2</sub>)<sub>n</sub>–  
37 wherein n = 3-6.

38 M, n, and p may be the same or different and are bridging groups of variable length from 3-12  
39 carbons, and

40 X is taken from a group consisting of nitrogen, sulfur, phosphorous and carbon.

1 4 The method of Claim one wherein said step of synthesizing further comprises the steps of:

- 2 -admixing an element taken from a group consisting of 2,4 dibromopropane and absolute  
3 ethanol into 1,2-diaminoethane hydrate;  
4 -heating the resulting mixture to approximately 50<sup>0</sup>C for about one hour;  
5 -adding potassium chloride;  
6 -continuing said heating for three hours;  
7 -filtering potassium bromide out of the mixture;  
8 -distilling the mixture at reduced pressure;  
9 -allowing the formation of top and bottom layers;  
10 -separating and distilling the top layer;  
11 -converting free amine in the distilled top layer to a tetrahydrochloride salt; and  
12 -converting said salt to a free amine by treatment with ammonium hydroxide.

1 5 The method of claim 4 wherein said step of converting to a tetrahydrochloride salt  
2 comprises adding hydrochloric acid to said distilled top layer.

1 6 The method of Claim 4 wherein said composition consists of 1,3-bis-[(2'-aminoethyl)-  
2 amino]propane and step of admixing a solution comprises preparing said solution by mixing  
3 1,3-dibromopropane and absolute ethanol in a ratio of approximately 1 to 3 per weight.

1 7 The method of Claim 6 wherein said step of admixing further comprises slowly adding said  
2 solution into 1,2-diaminoethane hydrate in a ratio of approximately 2.6 to 1 per weight.

1 8 The method of claim 7 wherein, the step of preparing said solution comprises mixing 15  
2 grams of 1,3-diaminopropane and 50 milliliters of absolute ethanol; and  
3 the step of slowly adding comprises adding said solution to 20 grams of potassium chloride;

1 9 The method of Claim 8 wherein said step of converting to a tetrahydrochloride salt  
2 comprises adding six molar concentration of hydrochloric acid.

1 10 The method of claim 2 wherein said step of selecting comprises:  
2 ascertaining the heats of formation of a set of said compounds; and choosing said compound in  
3 consideration of its heat of formation compared to the heats of formation of other compounds  
4 in said set.

1 11 The method of claim 10 wherein: said step of ascertaining comprises: calculating the heats  
2 at the formation of said set of compounds from their respective constituent atoms.

1 12 The method of claim 11 wherein said step of choosing comprises determining the  
2 stabilities of said set of compounds as a function of their respective heats of formation;  
3 wherein said stabilities are determined in inverse proportion to said respective heats of  
4 formation; and  
5 whereby the relative stabilities of the set of compounds are deemed indicative of ability to  
6 yield the most stable complex when reacted with a group of metals.

1 13 The method of Claim 12 wherein;  
2 said group of metals includes copper, cobalt, iron, zinc, cadmium, manganese and chromium.

1 14 The method of Claim 13 wherein said degenerative diseases comprise neurodegenerative  
2 diseases characterized by excess iron pools and said compound is selected from a group  
3 consisting of 2,2,2-piperidine and 2,3,2 adamantane.

1 15 The method of Claim 13 wherein said degenerative diseases comprise ischemic damage  
2 and pump failure post myocardial infarction characterized by iron-induced toxic redox effects  
3 and depletion of tissue zinc stores; and said compound is selected from a group consisting of  
4 zinc cyclam methylated, zinc cyclam adamantane, cyclam methylated and cyclam adamantane.


1 16 The method of Claim 13 wherein said degenerative diseases comprise neurodegenerative  
2 diseases and strokes; and said composition is selected from a group consisting of compositions  
3 having open ring metal binding molecules taken from a group consisting of compositions  
4 having copper binding molecules and manganese binding molecules.

1 17 The method of Claim 16 wherein said compositions having copper-binding molecules  
2 include 2,3,2 isopropyl on N1/N4; and  
3 said compositions having manganese-binding molecules include 3,3,3 tetramine.

1 18 The method of claim 13 wherein said degenerative diseases comprise neurodegenerative  
2 disorders, stroke, glaucoma, atherosclerosis, cardiomyopathy, ischemia, optic neuropathy,  
3 peripheral neuropathy, presbycusis and <sup>B</sup>cancer; and said composition is selected from  
4 derivatives of those compounds having the largest ring molecules.

1 19 The method of claim 18 wherein said compounds having the largest ring molecules  
2 includes 3,3,3 tetramine, cyclam adamantanes, cyclam 3,3,3 and compounds having alkyl  
3 substituted molecules.

1 20 The method of Claim 13 wherein said degenerative diseases comprise Parkinson's, Lou  
2 Gehrig's, Binswanger's, and Lewy Body diseases, Olivopontine Cerebellar Degeneration,  
3 stroke, glaucoma and optic neuropathy; and  
4 said composition is selected from a group of compositions having alkyl side chains.

1 21 The method of Claim 13 wherein said degenerative diseases comprise neurodegenerative  
2 diseases, ischemia post myocardial infarction and atherosclerosis; and  
3 said composition is selected from derivatives of compounds from a group consisting of  
4 piperidine, piperazine and adamantane 

1 22 The method of claim 3 wherein said degenerative diseases comprise stroke, diabetic  
2 neuropathy, peripheral neuropathy, Alzheimer's disease, atherosclerosis, ischemia, diabetes,  
3 presbycusis, cardiomyopathy and congestive heart failure; and said composition is derived  
4 from compounds having terminal nitrogen added molecule substitution with elements selected  
5 from a group consisting of glutathione, uric acid, ascorbic acid, taurine, estrogen,  
6 dehydroepiandrosterone, probucol, vitamin E, hydroxytoluene, carvidilol,  $\alpha$  lipoic acid,  
7 tocopherols, ubiquinone, phylloquinone, carotenes, menadione, glutamate, succinate, acetyl-l-  
8 carnitine, co-enzyme Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone,  
9 idebenone, dantrolene and phosphorous.



1 23 The method of Claim 22 wherein said degenerative disease comprises stroke; and said  
2 composition consists of uric acid polyamine.

1 24 The method of Claim 22 wherein said <sup>B</sup>degenerative disease comprises diabetes; and said  
2 composition is derived from compounds selected from a group consisting of phosphorous,  
3 taurine, CoEnzyme Q,  $\alpha$  lipoic acid, tocopherol, succinate, glutamate and acetyl-l-carnitine  
4 polyamines.

1 25 The method of Claim 22 wherein said degenerative disease comprises Alzheimer's disease  
2 and presbycusis; and  
3 said composition is derived from compounds selected from a group consisting of  $\alpha$  lipoic acid  
4 and acetyl-l-carnitine polyamines.

1 26 The method of Claim 22 wherein said degenerative disease comprises atherosclerosis; and  
2 said composition selected from a group consisting of tocopherol polyamine and coenzyme Q  
3 polyamine.

1 27 The method of Claim 22 wherein said degenerative disease comprises ischemia  
2 and  
3 said composition is selected from a group consisting of tocopherol polyamine and coenzyme Q  
4 polyamine.

1 28 The method of Claim 22 wherein said diseases comprise myocardial degeneration and  
2 congestive heart failure; and said composition consists of coenzyme Q polyamine.

1 29. The method of Claim 22 wherein said degenerative diseases comprise cancer; and  
2 said composition is taken from a group consisting of cobalt di-homocysteine  
3 polyamines.

1 30 The method of Claim 2 wherein said step of converting comprises adjusting the in vivo  
2 half life and pharmacokinetic properties of said composition by selective terminal nitrogen  
3 substitutions.

1 31 The method of Claim 2 wherein said step of converting comprises adjusting the in vivo  
2 half life and pharmacokinetic properties of said composition by addition of side chains on  
3 amino or methylene groups. *B*

1 32 The method of Claim 2 wherein said step of selecting comprises:  
2 finding the octanol / water coefficients of partition of a series of said compounds; and  
3 picking said compound in consideration of its octanol / water coefficient compared to the  
4 octanol water coefficients of other compounds in said series.

1 33 The method of Claim 32 wherein said step of picking comprises determining the abilities  
2 of said series of compounds to pass through the intestinal, blood brain and blood retinal  
3 barriers as a function of their respective octanol / water coefficients; wherein said abilities are  
4 determined according to a distribution curve centered about 2 and having a useful range  
5 extending towards 0.5 and 4, the numbers being log values.

1 34 The method of Claim 2 wherein said step of selecting comprises;

1 34 The method of Claim 2 wherein said step of selecting comprises;  
2 measuring pKas of a list of said compounds; and  
3 selecting said compound in consideration of its pKas compared to the pKa's of other  
4 compounds on the list.

1 35 The method of Claim 34 wherein said step of selecting comprises;  
2 selecting a composition with higher pKas in the treatment a disease characterized by lower  
3 tissue pH.

1 36 The method of Claim 35 wherein said diseases include ischemia post myocardial infarction  
2 and diabetic ketoacidosis.

1 37 The method of Claim 2 wherein said step of selecting comprises determining the respective  
2 likely efficiency of said compounds in consideration of the disease target to be treated and the  
3 route of administration.

1 38 The method of Claim 20 wherein;  
2 said compound consisting of pyridine tetramine.

1 39 The method of Claim 20 wherein said degenerative disease consists of Alzheimer's  
2 disease; and  
3 said compound comprises acetyl-l-carnitine polyamine.

1 40 The method of Claim 22 wherein said degenerative disease consists of diabetes; and

2 said compounds are selected from a group consisting of 2,3,2 piperidine, glutamate polyamine,  
3 succinate polyamine, chromium tetramine and vanadyl tetramine and phosphorous polyamine.

1 41 The method of Claim 2 wherein said degenerative diseases comprise peripheral and optic  
2 neuropathy; and

3 said compounds comprise taurine polyamine and  $\alpha$  lipoic acid polyamines.

1 42 The method of Claim 2 wherein said degenerative diseases comprise glaucoma; and said  
2 compounds comprise adamantane 2,3,2 tetramine and adamantane cyclam.

1 43 The method of Claim 3 wherein said degenerative disease comprise presbycusis; and said  
2 compounds comprise  $\alpha$  lipoic acid polyamine and acetyl-l-carnitine polyamine.

1 44 The method of Claim 4 wherein said composition consists of (2-aminoethyl){3-[(2-  
2 aminoethyl)amino]-1-methylbutyl}amine; and said step of admixing a solution comprises  
3 preparing said solution by mixing 2,4 dibromopropane and absolute ethanol in a ratio of  
4 approximately 1 to 20 per weight.

1 45 The method of claim 44 wherein said step of admixing comprises slowly adding said  
2 solution into 1,2-diaminoethane hydrate in a ratio of approximately 44 to 1 per weight.

1 46 The method of claim 45 wherein said step of converting to a tetrahydrochloride salt  
2 comprises of adding hydrochloric acid.

1 47 The method of Claim 2 wherein said composition consists of (2-aminoethyl){3-[(2-  
2 aminoethyl)amino]-1-methylbutyl}amine; and

3 said step of synthesizing further comprises; the steps of  
4 -admixing a solution of an element, taken from a group consisting of 1,3-diaminopropane and  
5 N,N-dimethyl-1,3-propanediamine and ethanol into 2-chloromethylpyridine in water;  
6 -adjusting the pH of the resulting mixture to 9 by addition of 10% sodium hydroxide;  
7 -stirring the mixture at room temperature and maintaining the pH between 8 and 9 by addition  
8 of sodium hydroxide over 3 days;  
9 -allowing solvents to evaporate; and  
10 -extracting residues with CH<sub>2</sub>Cl<sub>2</sub>.

1 48 The method of Claim 47 wherein said step of admixing a solution further comprises adding  
2 said solution into chloromethyl pyridine in water in a ratio of approximately 5 to 3 per weight  
3 wherein said chloromethylpyridine is diluted into water in a ratio of approximately 1 to 5 per  
4 weight.

1 49 The method of claim 48 wherein said step of admixing a solution comprises preparing said  
2 solution in a ratio of approximately 1 to 50 per weight.

1 50 The method of Claim 49 wherein said steps of synthesizing comprises synthesizing  
2 (2-pyridylmethyl){3-[(2-pyridylmethyl)amino]propyl}amine; and  
3 said step of admixing a solution further comprises preparing said solution by mixing 1,3-  
4 diaminopropane in water with ethanol.

1 51 The method of claim 50 when said step of synthesizing further comprises synthesizing  
2 methyl(3-[methyl(2-pyridylmethyl)amino]propyl)(2-pyridylmethyl)amine; and said step of  
3 admixing a solution further comprises preparing said solution by mixing N,N-dimethyl-1,3  
4 propanediamine in water with ethanol.

1 52 The method of claim 2 wherein said step of synthesizing comprises the steps of a  
2 preparation by adding a first solution of 1,3 diaminopropane and absolute ethanol dropwise  
3 into a second solution of ethanol and an element taken from a group consisting of 1-  
4 (2chloroethyl)piperidine and 1-(2-chloroethylpiperizine) and admixing over approximately 30  
5 minutes;  
6 stirring said preparation over approximately 24 hours;  
7 evaporating the solvents in said preparation;  
8 extracting the residue using a volume of  $\text{CH}_2\text{Cl}_2$  dried over  $\text{Na}_2\text{SO}_4$  and evaporated to dryness;  
9 purifying the resulting composition by converting to its hydrochloride salt by adding  
10 hydrochloric acid; and  
11 converting said salt to its free amine by treatment with  $\text{NH}_4\text{OH}$ .

1 53 The method of claim 52 wherein said step of mixing a preparation comprises forming said  
2 first solution of 1,3 diaminopropane and ethanol in a ratio of approximately 1 to 100 per  
3 weight and adding said first solution into said second solution in a ratio of approximately 1 to  
4 1 by weight.

1 54 The method of Claim 2 wherein said composition consists of

2 [2-(methylethylamino)ethyl](3-{[2-(methylamino)ethyl]amino}propyl)amine; and said step of

3 synthesizing further comprises; preparing of first mixture of magnesium turnings,

4 1,3-bis-[(2'-aminoethyl)-amino]propane, benzene and acetyl chloride respective approximate  
5 percentage of 0.6%, 8.5%, 8.45%, and 6.4% per weight;

6 cooling said first mixture;

7 separating the mixture into a liquid phase and a solid phase;

8 preparing a second mixture by mixing said solid phase with ether;

9 preparing a solution by pouring said second mixture over ice;

10 preparing a third mixture by adding said solution to said liquid phase;

11 washing said third mixture with sodium bicarbonate;

12 washing said third mixture with water.

1 55 The method of Claim 2 wherein said step of synthesizing comprises converting the starting

2 di - or tetramine component, at least one of said components in said compounds to the

3 corresponding N-substituted compound by treatment with an alkyl halide; and

4 purifying said composition by conversion to a salt through addition of hydrochloric acid.

1 56 The method of Claim 2 wherein said composition consists of (2-aminoethyl){3-[(2-  
2 aminoethyl)methylamino]propyl}methylamine, and

3 said step of synthesizing further comprises:

4 preparing a first solution of N,N-dimethyl-1,3-propanediamine and ethanol in a ratio of  
5 approximately 1 to 50 per weight;

6 preparing a second solution of 2-chloroethylamine and ethanol in a ratio of approximately 1 to  
7 17 per weight;  
8 combining said first and second solutions into a third solution;  
9 stirring said third solution at room temperature for approximately 20 hours;  
10 evaporating solvents in said third solution; and  
11 extracting residues in said solution with a volume of  $\text{CH}_2\text{Cl}_2$ .

1 57 The method of Claim 2 wherein said composition consists of  
2 [2-(bicyclo[3.3.1]non-3-ylamino)ethyl](3-{2-(bicyclo[3.3.1]non-3-  
3 ylamino)ethyl}amino)propyl)amine, and said step of synthesizing further comprises heating  
4 for approximately 6 hours at  $215^\circ\text{C}$  a mixture of 1-bromoadamantane and 2,3,2-tetramine in a  
5 mol ratio of approximately 1 to 5;  
6 admixing said mixture into a solution of  $2\text{NHCl}$  and ether having a ratio of approximately 1.25  
7 to 1 per weight, in a ratio of approximately 1 to 9 per weight;  
8 separating the aqueous layer and alkalinizing said layer in a volume of 50% aqueous  $\text{NaOH}$ ;  
9 extracting with ether;  
10 drying the extract over  $\text{K}_2\text{CO}_3$ ; and  
11 evaporating to an oil.

1 58 The method of Claim 2 wherein said composition consists of [2-  
2 (methylethylamino)ethyl](3 {[2-(methylamino)ethyl]amino}propyl)amine; and  
3 said methylating step of synthesizing further comprises;  
4 methylating terminal nitrogens of 2,3,2 tetramine by refluxing in the presence of benzene and  
5 acetyl chloride.



59 The method of Claim 58 wherein said step of synthesizing further comprises;  
preparing a first mixture of magnesium turnings;  
of 1,3-bis-[(2'-aminoethyl)-amino]propane, benzene and acetyl chloride respective  
approximate percentage of 0.6%, 8.5%, 8.45%, and 6.4% per weight;  
cooling said first mixture;  
separating the mixture into a liquid phase and a solid phase;  
preparing a second mixture by mixing said solid phase with ether;  
preparing a solution by pouring said second mixture over ice;  
preparing a third mixture by adding said solution to said liquid phase;  
washing said third mixture with sodium bicarbonate;  
washing said third mixture with water;  
drying said third mixture over  $\text{CaCl}_2$ ;  
filtering said third mixture;  
preparing a fourth mixture of said third mixture sodium hydride and N,N,-dimethylformamide  
in a ratio of approximately 2.5, 1 and 37.5 respectively per weight;  
heating said fourth mixture under  $\text{N}_2$  at approximately  $60^\circ\text{C}$  for about three hours;  
treating said fourth mixture with approximately  $\frac{1}{4}$  its volume of iodomethane;  
stirring said treated fourth mixture at  $50^\circ\text{C}$  for approximately 24 hours;  
quenching said treated fourth mixture with 95% ethanol;  
removing volatiles at reduced pressure;  
watering with addition of approximately  $\frac{1}{2}$  volume of water;  
extracting organic products with approximately three  $\frac{1}{2}$  volumes of chloroform;  
washing said organic products with water and  $\text{NaCl}$ ;  
drying said organic products over anhydrous sodium sulfate;

concentrating into an oil;  
purifying said oil by flash chromatography with ¼ hexanes-ethyl acetate as eluent into an acetylated oil of said composition;  
forming a solution of said acetylated oil, potassium hydroxide, methanol and water in respective proportions of 1, 3, 23 and 5 per weight respectively;  
heating said solution under reflux for about 24 hours;  
removing methanol at reduced pressure;  
extracting into ether;  
washing with NaCl;  
drying over sodium sulfate;  
concentrating under vacuum;  
purifying by flash chromatography; and  
evaporating solvents.

60 The method of Claim 2 wherein said composition consists of [2-(dimethylamino)ethyl](3-  
{[2-(dimethylamino)ethyl]methylamino}propyl)methylamine; and  
said steps of synthesizing further comprises;  
refluxing for about 20 hours a solution of 2,3,2 tetramine, formic acid and 37% formaldehyde  
and water in a weight proportions of approximately 1,10,10 and 1 respectively;  
evaporating solvents from said solution;  
making said solution basic by addition of NaOH; and  
extracting residues with 3 times 1½ volume of CH<sub>2</sub>Cl<sub>2</sub>.

1 61 The method of Claim 2 wherein said composition consists of 2-[3-(2-  
2 aminoethylthio)propylthio]ethylamine; and  
3 said step of synthesizing further comprises:  
4 preparing a first solution of 1,3-dimercaptopropane and water in a weight ration of about 1 to  
5 50;  
6 preparing a second solution of NaOH and water in a weight ratio of about 1.5 to 10;  
7 forming a first mixture by mixing said first and second solutions in a weight ratio of about 5 to  
8 1;  
9 forming a third solution of 2-chloroethylamine and ethanol in a weight ratio of about 8.5 to 1;  
10 admixing said solution into said mixture in a ratio of about 1 to 3.8;  
11 refluxing said mixture over approximately 8 hours;  
12 evaporating solvents from said refluxed mixture;  
13 extracting residues with CH<sub>2</sub>Cl<sub>2</sub>.

1 62 The method of Claim 2 wherein said composition consists of:  
2 1,4,8,11-tetraaza-1,4,8,11-tetramethylcyclotetradecane; and  
3 said steps of synthesizing comprises:  
4 refluxing for about 18 hours a solution of cyclam, formic acid, 37% formaldehyde and water in  
5 weight proportions of approximately 3.3, 4.5 and 1 respectively;  
6 adding water to said solution in a weight ratio of approximately 0.5 to 1;  
7 cooling said solution to about 5°C;  
8 adjust the pH of said solution to above 12 with NaOH;  
9 extracting the solution with CH<sub>2</sub>Cl<sub>2</sub>;

1 63 The method of Claim 2 wherein said composition consists of 1,4,8,11-tetraaza-1,4,8,11-  
2 tetra(2-piperidylethyl)cyclotetradecane; and said step of synthesizing further comprises:  
3 preparing a first solution of cyclam and  $\text{CH}_2\text{Cl}_2$  in a weight ratio of approximately 1 to 50;  
4 preparing a second solution of NaOH and water in a weight ratio of approximately 1 to 31;  
5 preparing a mixture of said first and second solution in a weight ratio of approximately 1 to 1;  
6 preparing a third solution of 1-(2-chloroethyl)piperidine and  $\text{CH}_2\text{Cl}_2$  in a weight ratio of  
7 approximately 1 to 14;  
8 adding said third solution dropwise into said mixture in a weight ratio of about 1 to 2;  
9 stirring said mixture over about 24 hours;  
10 evaporating solvents; and  
11 extracting residues with  $\text{CH}_2\text{Cl}_2$ .

1 64 The method of Claim 2 wherein said composition consists of 1,4,8,11-tetraaza-1,4,8,11 -  
2 tetrabicyclo[3.3.1]non-3-ylcyclotetradecane; and  
3 said step of synthesizing further comprises:  
4 forming a first solution of cyclam and ethanol in a weight ratio of approximately 1 to 100;  
5 forming a second solution of 1-bromoadamantane and ethanol in a weight ratio of 1 to 23;  
6 forming a mixture by adding said second solution dropwise into said first solution in a weight  
7 ratio of about 1 to 1, over 30 minutes;  
8 heating said mixture to reflux over about 20 hours;  
9 evaporating said solution under reduced pressure; and  
10 extracting residue from said solution with  $\text{CH}_2\text{Cl}_2$ ;

1 65 The method of Claim 2 wherein said composition consists of

2 1,4,8,11-tetraaza-1,4,8,11-tetraethylcyclotetradecane; and

3 said step of synthesizing further comprises:

4 forming a solution of cyclam and DMF in a weight ratio of approximately 1 to 50;

5 admixing under stirring small portions of NaH in a weight ratio of about 1 to 12.5;

6 heating said solution for about three hours at about 60°C;

7 admixing iodoethane in a single portion into said solution in a weight ratio of about 1 to 17.5;

8 heating said solution at about 60°C over about 18 hours;

9 quenching the solution with about 95% ethanol;

10 extracting residue with CH<sub>2</sub>Cl<sub>2</sub>.

1 66 The method of Claim 2 wherein said composition consists of N,N'-(2'  
2 dimethylphosphinoethyl)-propylenediamine; and the step of synthesizing further comprises:  
3 incorporating phosphorus into a molecule of propylenediamine in place of two of its nitrogen  
4 atoms by addition and reduction reactions.

1 67 The method of Claim 66 wherein said step of incorporating comprises:

2 preparing a first solution by dissolving propylenediamine into ethanol in a weight ratio of  
3 about 1 to 50;

4 admixing dimethylvinylphosphine sulfide into said solution in a weight ratio of about 1 to 22;

5 heating at reflux said solution for about 72 hours;

6 evaporating solvents under reduced pressure, leaving a residue.

1 68 The method of Claim 67 wherein said step of incorporating further comprises:

2 dissolving said residue in chloroform;

3 washing said residue with NaOH; and

4 drying said residue over  $\text{MgSO}_4$ .

1 69 The method of Claim 68 wherein said step of synthesizing further comprises:

2 removing solvents in said residue under reduced pressure to yield an oil,

3 crystallizing said oil with ethyl acetate;

4 preparing a suspension of  $\text{LiAlH}_4$  in dry dioxane in a weight ratio of about 1 to 100;

5 admixing said oil into said suspension;

6 to yield a mixture;

7 refluxing said mixture for about 36 hours;

8 cooling said mixture; and

9 adding a solution of dioxane in water and NaOH into said mixture.

1 70 The method of Claim 2 wherein said diseases consist of diabetes and abnormal low density

2 lipoprotein (LDL) to high density lipoprotein (HDL) ratio and said composition is selected

3 from a group consisting of vanadyl 2,3,2-tetramine and chromium 2,3,2-tetramine; and

4 said step of synthesizing further comprises reacting a metallic salt with 2,3,2-tetramine in an  
5 ethanol solution.

1 71 The method of Claim 70 wherein said step of reacting comprises:

2 forming a first solution of 2,3,2 tetramine in ethanol in a weight ratio of about 1 to 20;

3 forming a second solution of vanadyl acetylacetonate in ethanol in a weight ratio of about 1 to  
4 275;

5 admixing said second solution into said first solution in a volume ratio of about 1 to 1; and

6 refluxing said solution for almost 30 minutes.

1 72 The method of Claim 70 wherein said step of reacting further comprises:

2 preparing a first solution of 2,3,2-tetramine in ethanol in a weight ratio of about 1 to 20;

3 preparing a second solution of chromium (III) nitrate in ethanol in a weight ratio of about 1 to

4 80;

5 admixing said second solution into said first solution in a volume ratio of about 1 to 1; and

6 refluxing said solution for about 30 minutes.

1 73 The method of Claim 55 wherein said step of converting comprises using amines to attach

2 alkyl halide in a nucleophilic substitution of N atoms.